

 $\tt TTTGGAAGCAGCCACTTCTGGTCCCCATCCCTGGAGCGATCGAGCGCAGGATCTGCTGTCCCAT$ $\tt GGGACAGCAGATCTCTCTCCCAGTGCACAGCTTCTCCTGCTGTTTTCCCTGCTTCCAGTG$ $\tt CTCCAGGTGGCCCAACCAGGCCAGGCACCCCAGGACCAGCCCCTGTGGACACTTTTGGAGCAGT$ ${\tt ACTGCCACAGGACCACAATTGGGAATTTTTCAGGTCCCTACACCTACTGCAACACGACCTTGGA}$ CCAGATCGGGACCTGCTGGCCACAGAGCGCACCCGGAGCCCTAGTAGAGAGACCGTGCCCCGAG TACTTCAATGGCATCAAGTACAACACGACCCGGAATGCCTACAGAGAGTGCCTGGAGAACGGGA $\tt CCTGGGCCTCAAGGGTCAACTACTCACACTGCGAACCCATTTTGGATGACAAGCAGAGAAAGTA$ TGACCTGCATTACCGAATCGCCCTCATTGTCAACTACCTGGGTCACTGTGTTTCCGTGGTGGCC $\tt CTGGTGGCCGCTTTCCTGCTTTTCCTAGTGCTGCGGAGTATCCGCTGCCTGAGGAATGTGATCC$ ACTGGAACCTCATCACCACCTTCATTCTGAGAAACATCGCGTGGTTCCTGCTGCAACTCATCGA $\tt CCACGAAGTGCACGAGGGCAATGAGGTCTGGTGCCGCTGCATCACCACCATCTTCAACTATTTT$ CGTACTCCACAGAGCACCTGCGCAAGTGGCTTTTCCTCTTCATTGGATGGTGCATTCCCTGCCC TATCATCATCGCCTGGGCAGTTGGCAAACTCTACTATGAGAATGAGCAGTGCTGGTTTGGCAAG GAAGCTGGTGATTTGGTGGACTACATCTACCAGGGCCCCGTCATGCTTGTGCTGTTGATCAATT TTGTATTTCTGTTTAACATCGTCAGGATCCTGATGACGAAGTTACGAGCATCCACCACGTCCGA GACAATCCAATACAGGAAGGCAGTGAAGGCCACGCTGGTCCTCCCCCTGTTGGGCATCACC ${\tt TACATGCTCTTTGTCAATCCTGGCGAGGACGACCTGTCCCAGATTGTGTTCATCTACTTCA}$ ${\tt ACTCTTTCCTGCAGTCCTTCCAGGGTTTCTTTGTGTCCGTTTTCTACTGCTTCTTCAATGGAGA}$ GGTGCGCGCGCCCTGAGAAAGCGGTGGCACTCGGGGCAGGACCACCACGCCCTCCGGGTGCCT GTGCGCCGGGCCATGTCCATCCCTACGTCGCCCACCAGGATCAGCTTCCACAGCATCAAGCAGA CAGCTGCTGTGACCCTCTGTCACCGTCTGCCCGGCAGTCCACCACTGAGGCAGCTTCTCCAT CCTTTACAGCCTTCCCCTGGGTCCTCCTTGCTACCCTGACCCACAGGTACAAGGTACAGGAGAA GGGAGGAGAACGAACACTCCC (SEQ ID NO:1)

FIGURE 1

underlined = deleted in targeting construct

[] = sequence flanking Neo insert in targeting construct

 $\tt CTCCTTTGGAAGCAGCCACTTCTGGTCCCCATCCCTGGAGCGATCGAGCGCAGGATCTGC$ TGTCCCATGGGACAGCAGATCTCTCTCCCAGTGCACAGCTTCTCCTCTGCCTGTTTTCC CTGCTTCCAGTGCTCCAGGTGGCCCAACCAGGCCAGGCACCCCAGGACCAGCCCCTGTGG ACACTTTTGGAGCAGTACTGCCACAGGACCACAATTGGGAATTTTTCAGGTCCCTACACC TACTGCAACACGACCTTGGACCAGATCGGGACCTGCTGGCCACAGAGCGCACCCGGAGCC CTAGTAGAGAGACCGTGCCCCGAGTACTTCAATGGCATCAAGTACAACACGACCC [GGAA $\tt TGCCTACAGAGAGTGCCTGGA[\underline{GAACGGGACCTGGGCCTCAAGGGTCAACTACTCACACT}$ GCGAACCCATTTTGGATGACAAGCAGAGAAAGTATGACCTGCATTACCGAATCGCCCTCA TTGTCAACTACCTGGGTCACTGTGTTTCCGTGGTGGCCCTGGTG [GCCGCTTTCCTGCTT TTCCTAGTGCTGCG] GAGTATCCGCTGCCTGAGGAATGTGATCCACTGGAACCTCATCAC CACCTTCATTCTGAGAAACATCGCGTGGTTCCTGCTGCAACTCATCGACCACGAAGTGCA CAACTTCTTCTGGATGTTTGTGGAGGGCTGCTACCTGCACACGGCCATTGTCATGACGTA TATCATCATCGCCTGGGCAGTTGGCAAACTCTACTATGAGAATGAGCAGTGCTGGTTTGG ${\tt CAAGGAAGCTGGTGATTTGGTGGACTACATCTACCAGGGCCCCGTCATGCTTGTGCTGTT}$ GATCAATTTTGTATTTCTGTTTAACATCGTCAGGATCCTGATGACGAAGTTACGAGCATC CCTGTTGGGCATCACCTACATGCTCTTCTTTGTCAATCCTGGCGAGGACGACCTGTCCCA GATTGTGTTCATCTACTTCAACTCTTTCCTGCAGTCCTTCCAGGGTTTCTTTGTGTCCGT TTTCTACTGCTTCTTCAATGGAGAGGTGCGCGCGCCCTGAGAAAGCGGTGGCACTCGGG GCAGGACCACCACGCCTCCGGGTGCCTGTGCGCCGGGCCATGTCCATCCCTACGTCGCC CACCAGGATCAGCTTCCACAGCATCAAGCAGACAGCTGCTGTGTGACCCTCTGTCACCGT CTGCCCGGCAGTCCACCACTGAGGCAGCTTCTCCATCCTTTACAGCCTTCCCCTGGGTCC

FIGURE 2A

Gene Sequence Structure 441 bp 582 bp Sequence Deleted Size of full-length cDNA: 1557 bp Targeting Vector* (genomic sequence) LacZ-Neo Construct Number: 3050 Cassette 5' arm 3' arm Arm Length: 5': 3.1 kb 3': 2.5 kb 5'>AGCCCTATGTGTAATTTTCAT 5'>GCCGCTTTCCTGCTTTTCCTA ATAAATGACTCATATTAGCTTTCA GTGCTGCGGTGAGTCCACCTCCAC GATATGCATTGTGTTTTCAGGTCT CCTGCTTCCTCCTTGTCTTTGCCT GGGAGAACTAAGGAGTGTGGACCT CTCCCAGACATTGTCTCTTCCATT - Targeting Vector TATCCTGCAGGTACTAGGGAGCCA CTGGGGCCCCGGGAACAGTAGCCA – - Endogenous Locus GGGAGGGCTTTTGAGGCGGGAGGG GAAGTGGGTTTAAGTCAGACCCCC CGTCCTGACTCTCAGTGGTTGGCA AGGGCCATGACCACCAGCCTGCCT TCTTCTCTAGGGAATGCCTACAGA GAAGGGTAGAGAGCAAGCCCAGCT * Not drawn to scale GGGACCACCAG<3'
(SEQ ID NO:3) GAGTGCCTGGA<3' (SEQ ID NO:2)

FIGURE 2B